

=> d his ful

(FILE 'HOME' ENTERED AT 16:35:41 ON 07 JUN 2006)

FILE 'REGISTRY' ENTERED AT 16:43:09 ON 07 JUN 2006

L1 STR
L2 50 SEA SSS SAM L1
L3 13161 SEA SSS FUL L1

FILE 'HCAPLUS' ENTERED AT 16:49:54 ON 07 JUN 2006

L4 846 SEA ABB=ON L3

FILE 'REGISTRY' ENTERED AT 16:51:04 ON 07 JUN 2006

E HISTONE DEACETYLASE/CN
L5 157 SEA ABB=ON HISTONE DEACETYLASE?/CN

FILE 'HCAPLUS' ENTERED AT 16:52:38 ON 07 JUN 2006

L6 13 SEA ABB=ON L4 AND (L5 OR ?HISTONE?(W)?DEACETYLASE?) (4A)?INHIBIT?

FILE 'REGISTRY' ENTERED AT 16:54:30 ON 07 JUN 2006

L7 STR
L8 0-SEA SSS SAM L7
L9 3 SEA SSS FUL L7
L10 13159 SEA ABB=ON L3 NOT L9

FILE 'HCAPLUS' ENTERED AT 16:59:25 ON 07 JUN 2006

L11 844 SEA ABB=ON L10
L12 13 SEA ABB=ON L11 AND (L5 OR ?HISTONE?(W)?DEACETYLASE?) (4A)?INHIBIT?

FILE 'HCAPLUS' ENTERED AT 17:01:19 ON 07 JUN 2006

L13 5 SEA ABB=ON L12 AND (PRD<20020723 OR PD<20020723) *5 cit's from CAPLUS*

FILE 'USPATFULL' ENTERED AT 17:02:10 ON 07 JUN 2006

L14 2 SEA ABB=ON L12 AND (PRD<20020723 OR PD<20020723) *2 cit's from USPatfull*

FILE HOME

FILE REGISTRY

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 6 JUN 2006 HIGHEST RN 887000-62-6

DICTIONARY FILE UPDATES: 6 JUN 2006 HIGHEST RN 887000-62-6

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH January 6, 2006

Please note that search-term pricing does apply when conducting SmartSELECT searches.

*
* The CA roles and document type information have been removed from *
* the IDE default display format and the ED field has been added, *
* effective March 20, 2005. A new display format, IDERL, is now *
* available and contains the CA role and document type information. *
*

Structure search iteration limits have been increased. See HELP SLIMITS for details.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

<http://www.cas.org/ONLINE/UG/regprops.html>

FILE HCAPLUS

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FILE COVERS 1907 - 7 Jun 2006 VOL 144 ISS 24
FILE LAST UPDATED: 6 Jun 2006 (20060606/ED)

New CAS Information Use Policies, enter HELP USAGETERMS for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

FILE USPATFULL

FILE COVERS 1971 TO PATENT PUBLICATION DATE: 6 Jun 2006 (20060606/PD)
FILE LAST UPDATED: 6 Jun 2006 (20060606/ED)
HIGHEST GRANTED PATENT NUMBER: US7058980
HIGHEST APPLICATION PUBLICATION NUMBER: US2006117448
CA INDEXING IS CURRENT THROUGH 6 Jun 2006 (20060606/UPCA)
ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 6 Jun 2006 (20060606/PD)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Feb 2006
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Feb 2006

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COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

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772.95

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE

TOTAL

ENTRY

SESSION

CA SUBSCRIBER PRICE

0.00

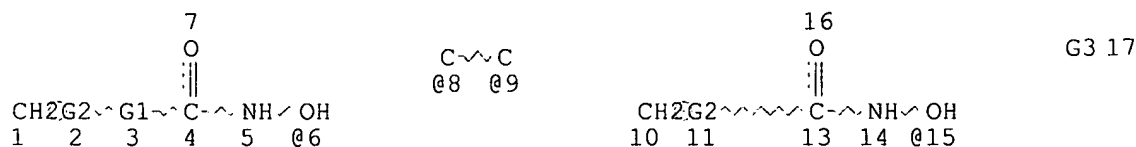
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SESSION WILL BE HELD FOR 60 MINUTES

STN INTERNATIONAL SESSION SUSPENDED AT 17:04:37 ON 07 JUN 2006

=> d que stat 113

L1 STR



VAR G1=C/CH/S/N/8-2 9-4

VAR G2=C/P

VAR G3=6/15

NODE ATTRIBUTES:

DEFAULT MLEVEL IS ATOM

DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

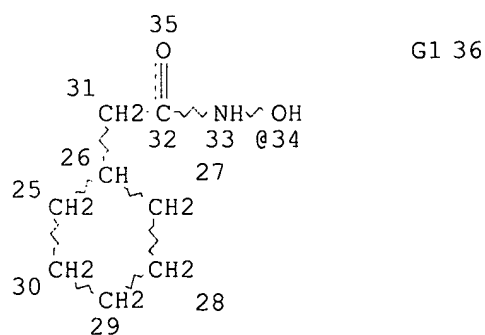
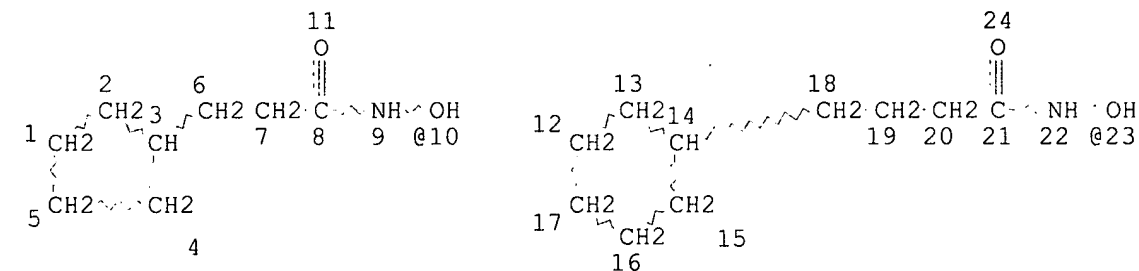
NUMBER OF NODES IS 16

STEREO ATTRIBUTES: NONE

L3 13161 SEA FILE=REGISTRY SSS FUL L1

L5 157 SEA FILE=REGISTRY ABB=ON HISTONE DEACETYLASE?/CN

L7 STR



VAR G1=10/23/34

NODE ATTRIBUTES:

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DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 36

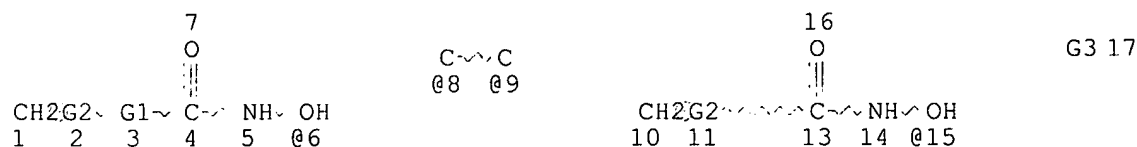
STEREO ATTRIBUTES: NONE

L9 3 SEA FILE=REGISTRY SSS FUL L7

L10 13159 SEA FILE=REGISTRY ABB=ON L3 NOT L9
 L11 844 SEA FILE=HCAPLUS ABB=ON L10
 L12 13 SEA FILE=HCAPLUS ABB=ON L11 AND (L5 OR ?HISTONE?(W)?DEACETYLAS
 E?) (4A)?INHIBIT?
 L13 5 SEA FILE=HCAPLUS ABB=ON L12 AND (PRD<20020723 OR PD<20020723)

=> d que stat l14

L1 STR



VAR G1=C/CH/S/N/8-2 9-4

VAR G2=C/P

VAR G3=6/15

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DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

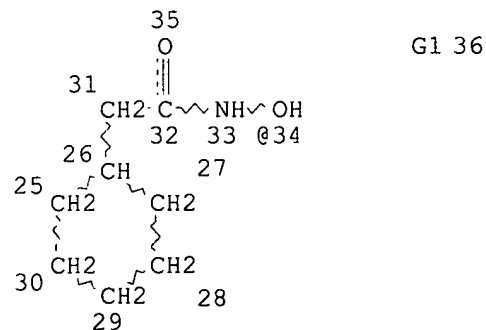
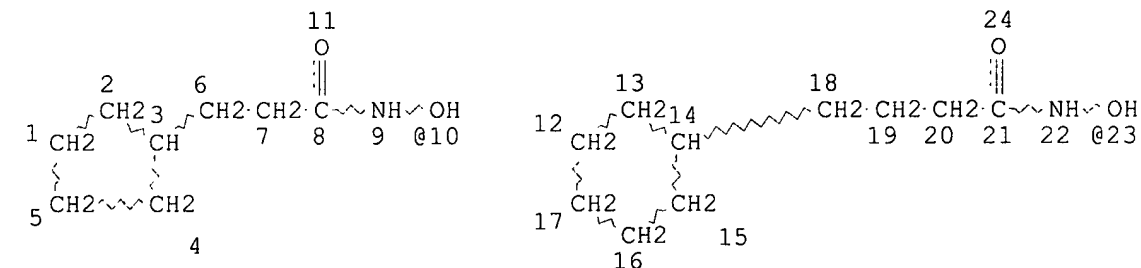
NUMBER OF NODES IS 16

STEREO ATTRIBUTES: NONE

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L5 157 SEA FILE=REGISTRY ABB=ON HISTONE DEACETYLASE?/CN

L7 STR



VAR G1=10/23/34

NODE ATTRIBUTES:

DEFAULT MLEVEL IS ATOM

DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 36

STEREO ATTRIBUTES: NONE

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L10 13159 SEA FILE=REGISTRY ABB=ON L3 NOT L9

L11 844 SEA FILE=HCAPLUS ABB=ON L10

L12 13 SEA FILE=HCAPLUS ABB=ON L11 AND (L5 OR ?HISTONE?(W)?DEACETYLAS
E?) (4A)?INHIBIT?

L14 2 SEA FILE=USPATFULL ABB=ON L12 AND (PRD<20020723 OR PD<20020723
)

=> d ibib abs hitstr 113 1-5

L13 ANSWER 1 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:737742 HCAPLUS

DOCUMENT NUMBER: 139:276884

TITLE: Preparation of sulfonyl-derivatives as novel inhibitors of histone deacetylase

INVENTOR(S): Van Emelen, Kristof; Arts, Janine; Backx, Leo Jacobus Jozef; De Winter, Hans Louis Jos; Van Brandt, Sven Franciscus Anna; Verdonck, Marc Gustaaf Celine; Meerpoel, Lieven; Pilatte, Isabelle Noeelle Constance; Poncelet, Virginie Sophie; Dyatkin, Alexey Borisovich

PATENT ASSIGNEE(S): Janssen Pharmaceutica N.V., Belg.; et al.

SOURCE: PCT Int. Appl., 139 pp.

CODEN: PIXXD2

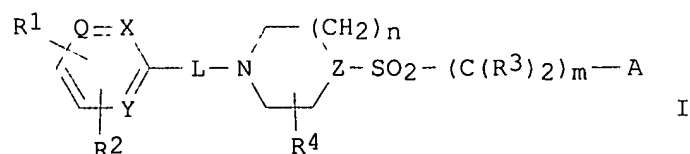
DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 8

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003076422	A1	20030918	WO 2003-EP2516	20030311 <--
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
CA 2476586	AA	20030918	CA 2003-2476586	20030311 <--
AU 2003218738	A1	20030922	AU 2003-218738	20030311 <--
EP 1485365	A1	20041215	EP 2003-711982	20030311 <--
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK			
BR 2003007575	A	20041221	BR 2003-7575	20030311 <--
US 2005113373	A1	20050526	US 2003-507708	20030311 <--
CN 1642931	A	20050720	CN 2003-805952	20030311 <--
JP 2005525380	T2	20050825	JP 2003-574641	20030311 <--
NZ 534830	A	20050826	NZ 2003-534830	20030311 <--
NO 2004004314	A	20041012	NO 2004-4314	20041012 <--
PRIORITY APPLN. INFO.:			US 2002-363799P	P 20020313 <--
			US 2002-420989P	P 20021024
			WO 2003-EP2516	W 20030311

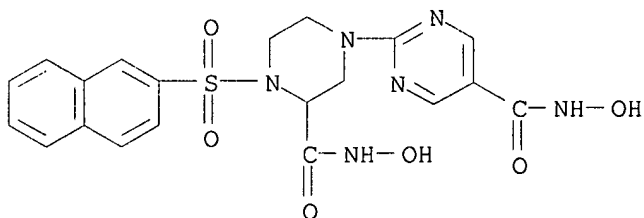
OTHER SOURCE(S): MARPAT 139:276884
GI

AB This invention comprises the novel compds. (I) (wherein n = 1-3, m = 1-4, Q, X, Y = N, CH; Z = N, CH; R1 = (un)substituted amido, acylamido, guandido, and other Zn chelating group, etc.; R2 = H, halo, OH, NH2, NO2, C1-6alkyl, C1-6alkoxy, CF3, di(C1-6alkyl)amino, HONH, naphthalenylsulfonylpyrazinyl; R3 = H aryl; R4 = H, HO, NH2, hydroxyC1-6alkyl, C1-6alkyl, C1-6alkoxy, arylC1-6alkyl, aminocarbonyl, hydroxycarbonyl, aminoC1-6alkyl, aminocarbonylC1-6alkyl, hydroxycarbonylC1-6alkyl, hydroxyaminocarbonyl, C1-6alkoxycarbonyl, C1-6alkylamino, di(C1-6alkyl)aminoC1-6alkyl; L = nul or bivalent radical selected from C1-6alkanediyl, amino, carbonyl or aminocarbonyl; A = aryl, cyclohexyl, heterocyclic derivs.), having histone deacetylase inhibiting enzymic activity; their preparation, compns. containing them and their use as a medicine. For example, 4-(4-(2-naphthylsulfonyl)piperazin-1-yl)-N-hydroxybenzamide in 100% yield was prepared by the hydrogenation of 4-(4-(2-naphthylsulfonyl)piperazin-1-yl)-N-(phenylmethoxy)benzamide (II) in THF by Pd/C as a catalyst. II was prepared from 1,1-dimethylethyl 4-(4-carboxyphenyl)-1-piperazinecarboxylate and O-(phenylmethyl)hydroxylamine hydrochloride in presence of dimethylaminopyridine in CH2Cl2 and diisopropylcarbodiimide, forming 1,1-dimethylethyl 4-[4-[(phenylmethoxy)amino]carbonylphenyl]-1-piperazinecarboxylate which was saponified and subsequently reacted with 2-naphthalenesulfonyl chloride to give the II.

IT 604769-20-2P
 RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (preparation of sulfonyl derivs. as histone deacetylase inhibitors and antitumor agent for treatment of cancer)

RN 604769-20-2 HCAPLUS

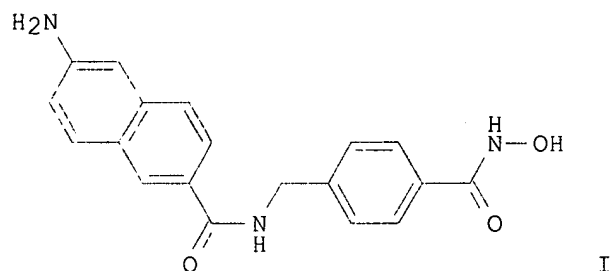
CN 5-Pyrimidinecarboxamide, N-hydroxy-2-[3-[(hydroxyamino)carbonyl]-4-(2-naphthalenylsulfonyl)-1-piperazinyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 2 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2003:678775 HCAPLUS
 DOCUMENT NUMBER: 139:214215
 TITLE: Preparation of N-hydroxycarboxamide derivatives as anticancer agents
 INVENTOR(S): Uesato, Shinichi; Nagaoka, Yasuo; Yamori, Takao
 PATENT ASSIGNEE(S): Osaka Industrial Promotion Organization, Japan
 SOURCE: PCT Int. Appl., 137 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003070691	A1	20030828	WO 2003-JP1681	20030218 <--
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
AU 2003211362	A1	20030909	AU 2003-211362	20030218 <--
PRIORITY APPLN. INFO.:			JP 2002-45310	A 20020221 <--
			JP 2002-235912	A 20020813
			WO 2003-JP1681	W 20030218
OTHER SOURCE(S):		MARPAT 139:214215		
GI				



AB The title N-hydroxycarboxamides with general formula of D-L2-B-N(R)-L1-A-CONHOH [wherein A = cycloalkylene, phenylene, naphthylene, anthrylene, phenanthrylene, cycloalkenylene, biphenylene, heterocycloalkylene, or heterocycloalkenylene, etc., with exclusions; B = CO, CS, NHCO, NHCS, SO₂, SO, S, O, CO₂, or OCO; D = cycloalkyl, adamantyl, Ph, naphthyl, anthryl, phenanthryl, cycloalkenyl, biphenyl, pyridyl, quinolyl, isoquinolyl, indolyl, heterocycloalkyl, or heterocycloalkenyl, etc., with exclusions; L1 and L2 = independently alkylene or none; R = H, alkyl, CHO, alkanoyl, PhCO, or PhCH₂CO] and tautomers, stereoisomers, or salts thereof are prepared as potent histone deacetylase (HDAC) inhibitors. The N-hydroxycarboxamide derivs. are useful in treating, relieving, and preventing diseases concerning cell proliferation. In particular, it is expected that these derivs. are highly efficacious as an anticancer agent or a carcinostatic agent. Moreover, it is expected that the above N-hydroxycarboxamide derivs. are efficacious as an immunosuppressant or a gene therapy potentiator and usable in treating, relieving, and preventing neurodegenerative diseases. For example, the compound I•HCl was prepared in a four-step synthesis in moderate yield. I showed IC₅₀ of 39 nM against human histone deacetylase (HDAC).

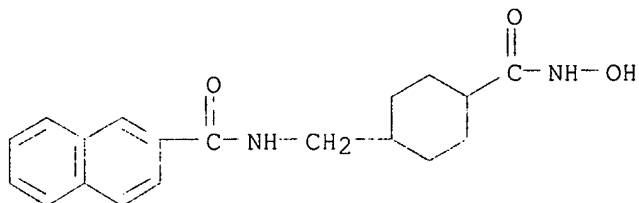
IT 471924-76-2P 471924-77-3P 471924-78-4P
 471924-84-2P 471924-85-3P 471924-88-6P
 471925-06-1P
 RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU

(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
(Uses)

(drug candidate; preparation of hydroxycarboxamide derivs. as anticancer agents)

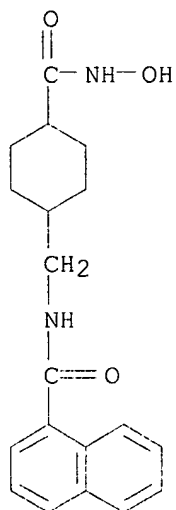
RN 471924-76-2 HCAPLUS

CN 2-Naphthalenecarboxamide, N-[[4-[(hydroxyamino)carbonyl]cyclohexyl]methyl]-
(9CI) (CA INDEX NAME)



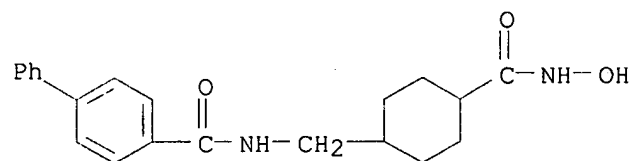
RN 471924-77-3 HCAPLUS

CN 1-Naphthalenecarboxamide, N-[[4-[(hydroxyamino)carbonyl]cyclohexyl]methyl]-
(9CI) (CA INDEX NAME)



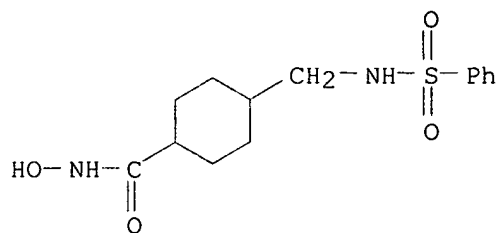
RN 471924-78-4 HCAPLUS

CN [1,1'-Biphenyl]-4-carboxamide, N-[[4-[(hydroxyamino)carbonyl]cyclohexyl]methyl]- (9CI) (CA INDEX NAME)



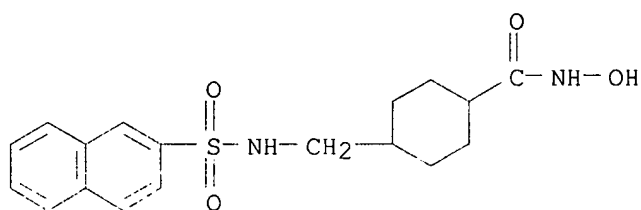
RN 471924-84-2 HCAPLUS

CN Cyclohexanecarboxamide, N-hydroxy-4-[[[(phenylsulfonyl)amino]methyl]- (9CI)
(CA INDEX NAME)



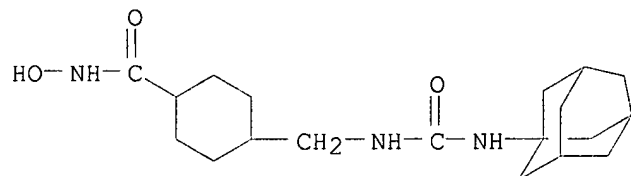
RN 471924-85-3 HCAPLUS

CN Cyclohexanecarboxamide, N-hydroxy-4-[[(2-naphthalenylsulfonyl)amino]methyl]- (9CI) (CA INDEX NAME)



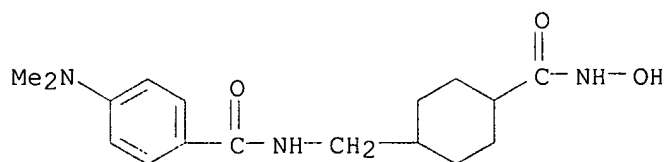
RN 471924-88-6 HCAPLUS

CN Cyclohexanecarboxamide, N-hydroxy-4-[[[(tricyclo[3.3.1.1.3,7]dec-1-ylamino)carbonyl]amino]methyl]- (9CI) (CA INDEX NAME)



RN 471925-06-1 HCAPLUS

CN Benzamide, 4-(dimethylamino)-N-[[4-[(hydroxyamino)carbonyl]cyclohexyl]methyl]- (9CI) (CA INDEX NAME)



IT 9076-57-7, Histone deacetylase

RL: BSU (Biological study, unclassified); BIOL (Biological study) (inhibitor; preparation of hydroxycarboxamide derivs. as anticancer agents)

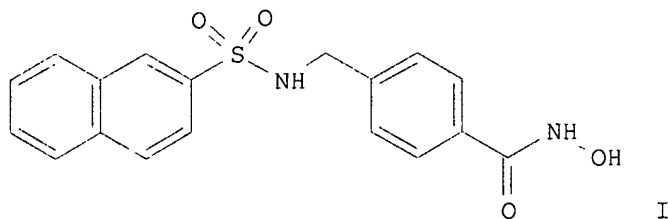
RN 9076-57-7 HCAPLUS

CN Deacetylase, histone (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

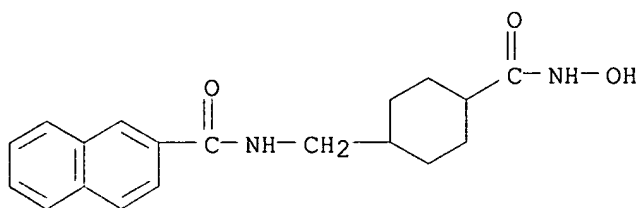
L13 ANSWER 3 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2002:324923 HCAPLUS
DOCUMENT NUMBER: 137:310681
TITLE: Novel histone deacetylase inhibitors: N-hydroxycarboxamides possessing a terminal bicyclic aryl group
AUTHOR(S): Uesato, Shinichi; Kitagawa, Manabu; Nagaoka, Yasuo; Maeda, Taishi; Kuwajima, Hiroshi; Yamori, Takao
CORPORATE SOURCE: Department of Biotechnology, Faculty of Engineering, Kansai University, Suita, Osaka, 564-8680, Japan
SOURCE: Bioorganic & Medicinal Chemistry Letters (2002), 12(10), 1347-1349
CODEN: BMCLE8; ISSN: 0960-894X
PUBLISHER: Elsevier Science Ltd.
DOCUMENT TYPE: Journal
LANGUAGE: English
OTHER SOURCE(S): CASREACT 137:310681
GI



AB Utilizing tranexamic acid as a starting material, a series of N-hydroxycarboxamides (e.g., I) were synthesized in order to seek new histone deacetylase (HDAC) inhibitors. Compound I showed antiproliferative activity against HDAC of IC₅₀ = 1100 nM. Further structure optimization involving the replacement of the 1,4-cyclohexylene group with the 1,4-phenylene group yielded the promising HDAC inhibitors which possess a terminal bicyclic aryl amide.

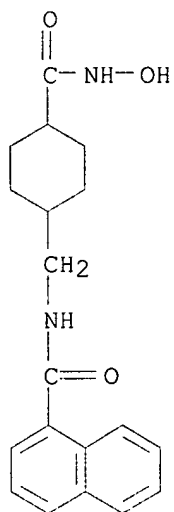
IT 471924-76-2P 471924-77-3P 471924-78-4P
471924-84-2P 471924-85-3P 471924-88-6P
471925-06-1P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)
(preparation of N-hydroxycarboxamides as antitumor agents)

RN 471924-76-2 HCAPLUS
CN 2-Naphthalenecarboxamide, N-[[4-[(hydroxyamino)carbonyl]cyclohexyl]methyl]-
(9CI) (CA INDEX NAME)



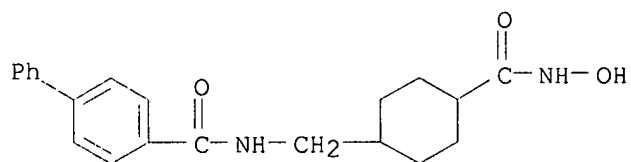
RN 471924-77-3 HCAPLUS

CN 1-Naphthalenecarboxamide, N-[[4-[(hydroxyamino)carbonyl]cyclohexyl]methyl]-
(9CI) (CA INDEX NAME)



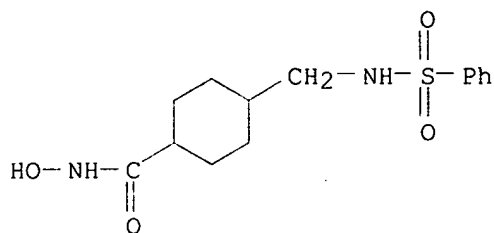
RN 471924-78-4 HCAPLUS

CN [1,1'-Biphenyl]-4-carboxamide, N-[[4-[(hydroxyamino)carbonyl]cyclohexyl]methyl]- (9CI) (CA INDEX NAME)



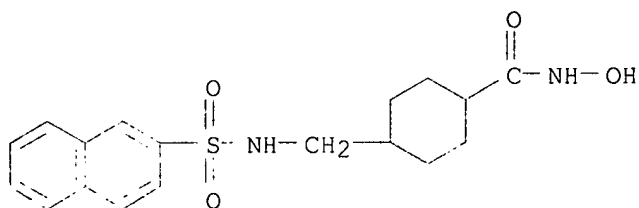
RN 471924-84-2 HCAPLUS

CN Cyclohexanecarboxamide, N-hydroxy-4-[[[(phenylsulfonyl)amino]methyl]- (9CI)
(CA INDEX NAME)



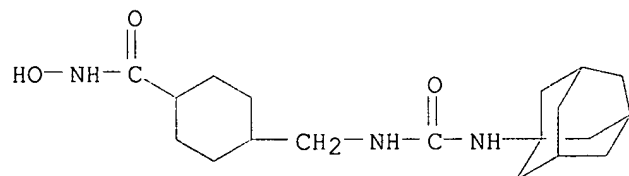
RN 471924-85-3 HCAPLUS

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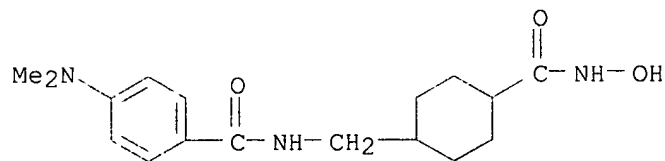
RN 471924-88-6 HCAPLUS

CN Cyclohexanecarboxamide, N-hydroxy-4-[[[(tricyclo[3.3.1.1.3,7]dec-1-ylamino)carbonyl]amino]methyl]- (9CI) (CA INDEX NAME)



RN 471925-06-1 HCAPLUS

CN Benzamide, 4-(dimethylamino)-N-[[4-[(hydroxyamino)carbonyl]cyclohexyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT:

11

THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 4 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:256222 HCAPLUS

DOCUMENT NUMBER: 136:294651

TITLE: Preparation of aryl-substituted N-hydroxy amides with amide linkages as HDAC inhibitors for treatment of

proliferative conditions

INVENTOR(S): Watkins, Clare J.; Romero-Martin, Maria-Rosario;
Moore, Kathryn G.; Ritchie, James; Finn, Paul W.;
Kalvinsh, Ivars; Loza, Einars; Starchenkov, Igor;
Dikovska, Klara; Bokaldere, Rasma Melita; Gailite,
Vija; Vorona, Maxim; Andrianov, Victor; Lolya, Daina;
Semenikhina, Valentina; Amolins, Andris; Harris, C.
John; Duffy, James E. S.

PATENT ASSIGNEE(S): Prolifix Limited, UK

SOURCE: PCT Int. Appl., 346 pp.
CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002026696	A1	20020404	WO 2001-GB4329	20010927 <--
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CA 2423868	AA	20020404	CA 2001-2423868	20010927 <--
AU 2001090134	A5	20020408	AU 2001-90134	20010927 <--
EP 1335898	A1	20030820	EP 2001-970014	20010927 <--
EP 1335898	B1	20051123		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
JP 2004509941	T2	20040402	JP 2002-531082	20010927 <--
EP 1598067	A1	20051123	EP 2005-15737	20010927 <--
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, CY, TR				
AT 310719	E	20051215	AT 2001-970014	20010927 <--
US 2004092598	A1	20040513	US 2003-381791	20030827 <--
PRIORITY APPLN. INFO.:				
			GB 2000-23985	A 20000929 <--
			US 2001-297785P	P 20010614 <--
			EP 2001-970014	A3 20010927 <--
			WO 2001-GB4329	W 20010927 <--

OTHER SOURCE(S): MARPAT 136:294651

AB The title compds. AQ1JQ2CONHOH [I; wherein A = aryl group; Q1 = aryl leader group having a backbone of at least 2 C atoms; J = NR1CO or CONR1; R1 = amido substituent; Q2 = acid leader group; and pharmaceutically acceptable salts, solvates, amides, esters, ethers, chemical protected forms, and prodrugs thereof] were prepared via solution phase and solid phase synthetic methods as histone deacetylase (HDAC) inhibitors for treatment of proliferative conditions, such as cancer and psoriasis. For example, 6-aminocaproic acid Me ester•HCl was coupled with 2-naphthoyl chloride in the presence of diisopropyl ethylamine in DMF to give the amide. Deesterification (79%), followed by conversion to the N-hydroxyamide using HONH2•HCl in the presence of 1,1'-carbonyldiimidazole in THF, afforded naphthalene-2-carboxylic acid (5-hydroxycarbamoylpentyl)amide II (PX105687) in 40% yield. The latter inhibited recombinant HDAC1 and HDAC2 with IC50 values of 33 nM and 29 nM, resp., and inhibited cell proliferation against the human cervical

adenocarcinoma (HeLa) cell line using cell proliferation reagent WST-1 with IC50 of 1.1 nM. Structure-activity relationship studies showed superior activity for I when (1) the backbone of Q1 had > 1 carbon atoms, and (2) the alkylene group Q2 had > 5 carbon atoms.

IT 408325-14-4P, PX 105552

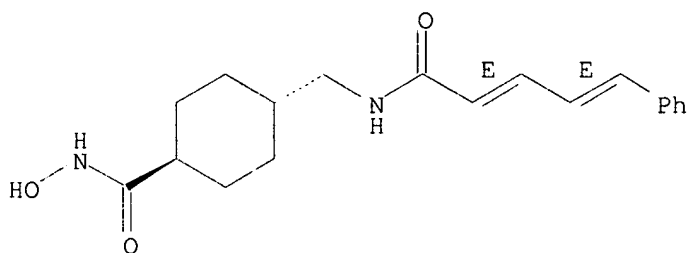
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(HDAC inhibitor; preparation of N-hydroxy amides with amide linkages as HDAC inhibitors for treatment of proliferative conditions)

RN 408325-14-4 HCAPLUS

CN Cyclohexanecarboxamide, N-hydroxy-4-[[[(2E,4E)-1-oxo-5-phenyl-2,4-pentadienyl]amino]methyl]-, trans- (9CI) (CA INDEX NAME)

Relative stereochemistry.
Double bond geometry as shown.



REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 5 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2001:396861 HCAPLUS

DOCUMENT NUMBER: 135:5455

TITLE: Preparation of hydroxamic acids as inhibitors of histone deacetylase

INVENTOR(S): Delorme, Daniel; Ruel, Rejean; Lavoie, Rico; Thibault, Carl; Abou-khalil, Elie

PATENT ASSIGNEE(S): Methylgene, Inc., Can.

SOURCE: PCT Int. Appl., 147 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

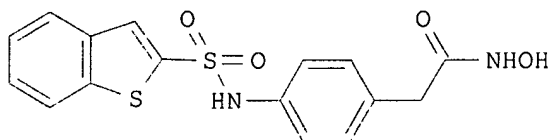
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001038322	A1	20010531	WO 2000-IB1881	20001122 <--
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
CA 2391952	AA	20010531	CA 2000-2391952	20001122 <--
EP 1233958	A1	20020828	EP 2000-981535	20001122 <--

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO, MK, CY, AL, TR

US 6541661	B1	20030401	US 2000-718265	20001122 <--
JP 2003514904	T2	20030422	JP 2001-540085	20001122 <--
AU 783504	B2	20051103	AU 2001-18768	20001122 <--
AU 2006200456	A1	20060302	AU 2006-200456	20060202 <--
PRIORITY APPLN. INFO.:			US 1999-167035P	P 19991123 <--
			AU 2001-18768	A3 20001122 <--
			WO 2000-IB1881	W 20001122 <--

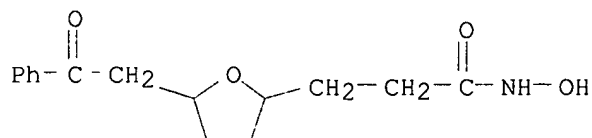
OTHER SOURCE(S): MARPAT 135:5455
 GI



AB The title compds. CyL1ArY1CONHZ [Cy = (un)substituted cycloalkyl, aryl, heteroaryl, etc.; L1 = (CH2)mW (wherein m = 0-4; W = CONH, SO2NH, NHCO, NHSO2, NHCONH); Ar = (un)substituted arylene which may be fused to an aryl, heteroaryl, etc.; Y1 = a bond, alkylene; Z = aniliny1, pyridyl, thiadiazolyl, OM (M = H, a pharmaceutically acceptable cation)], useful for inhibiting histone deacetylase enzymic activity, were prepared E.g., a multi-step synthesis of the title compound I which showed IC50 of 7 μ M against histone deacetylase in nuclear exts. from H446 cells (pooled HDACs), was given. The invention also provides compns. and methods for treating cell proliferative diseases and conditions.

IT 342373-15-3P
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (preparation of hydroxamic acids as inhibitors of histone deacetylase)

RN 342373-15-3 HCAPLUS
 CN 2-Furanpropanamide, tetrahydro-N-hydroxy-5-(2-oxo-2-phenylethyl)- (9CI)
 (CA INDEX NAME)



REFERENCE COUNT: 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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=> d ibib abs hitstr 114 1-2

L14 ANSWER 1 OF 2 USPATFULL on STN

ACCESSION NUMBER: 2004:121182 USPATFULL

TITLE: Carbamic acid compounds comprising an amide linkage as
hdac inhibitorsINVENTOR(S): Watkins, Clare J., Oxon, UNITED KINGDOM
Romero-Martin, Maria Rosario, Oxon, UNITED KINGDOM
Moore, Kathryn G., Oxon, UNITED KINGDOM
Ritchie, James, Oxon, UNITED KINGDOM
Finn, Paul W., Oxon, UNITED KINGDOM
Kalvinsh, Ivars, Riga, LATVIA
Loza, Einars, Riga, LATVIA
Starchenkov, Igor, Riga, LATVIA
Dikovska, Klara, Riga, LATVIA
Bokaldere, Rasma, Riga, LATVIA
Gailite, Vija, Riga, LATVIA
Vorona, Maxim, Riga, LATVIA
Andrianov, Victor, Riga, LATVIA
Lolya, Daina, Riga, LATVIA
Seminkhina, Valentina, Riga, LATVIA
Amolins, Andris, Riga, LATVIA
Harris, C.John, Kent, UNITED KINGDOM
Duffy, James E. S., Kent, UNITED KINGDOM

	NUMBER	KIND	DATE
PATENT INFORMATION:	US 2004092598	A1	20040513
APPLICATION INFO.:	US 2003-381791	A1	20030827 (10)
	WO 2001-GB4329		20010927

	NUMBER	DATE
PRIORITY INFORMATION:	GB 2000-23985	20000929 <--
DOCUMENT TYPE:	Utility	
FILE SEGMENT:	APPLICATION	
LEGAL REPRESENTATIVE:	Nixon & Varderhye, 8th Floor, 1100 North Glebe Rd, Arlington, VA, 22201-4714	
NUMBER OF CLAIMS:	188	
EXEMPLARY CLAIM:	1	
LINE COUNT:	9591	

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB This invention pertains to certain active carbamic acid compounds which inhibit HDAC activity and which have the formula (1) wherein: A is an aryl group; Q1 is an aryl leader group having a backbone of at least 2 carbon atoms; J is an amide linkage selected from: --NR1C(.dbd.O)--and --C(.dbd.O)NR1--; R1 is an amido substituent; and, Q2 is an acid leader group; and pharmaceutically acceptable salts, solvates, amides, esters, ethers, chemically protected forms, and prodrugs thereof. The present invention also pertains to pharmaceutical compositions comprising such compounds, and the use of such compounds and compositions, both in vitro and in vivo, to inhibit HDAC, and, e.g., to inhibit proliferative conditions, such as cancer and psoriasis.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 408325-14-4P, PX 105552

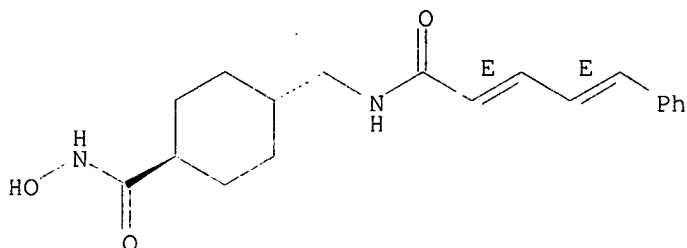
(HDAC inhibitor; preparation of N-hydroxy amides with amide linkages as HDAC

inhibitors for treatment of proliferative conditions)

RN 408325-14-4 USPATFULL

CN Cyclohexanecarboxamide, N-hydroxy-4-[[[(2E,4E)-1-oxo-5-phenyl-2,4-pentadienyl]amino]methyl]-, trans- (9CI) (CA INDEX NAME)

Relative stereochemistry.
Double bond geometry as shown.



L14 ANSWER 2 OF 2 USPATFULL on STN

ACCESSION NUMBER: 2003:89492 USPATFULL

TITLE: Inhibitors of histone deacetylase

INVENTOR(S): Delorme, Daniel, St. Lazare, CANADA
Ruel, Rejean, St. Lazare, CANADA
Lavoie, Rico, Lachine, CANADA
Thibault, Carl, Mascouche, CANADA
Abou-Khalil, Elie, Laval, CANADA

PATENT ASSIGNEE(S): MethylGene, Inc., Montreal, CANADA (non-U.S. corporation)

	NUMBER	KIND	DATE
PATENT INFORMATION:	US 6541661	B1	20030401
APPLICATION INFO.:	US 2000-718265		20001122 (9)

	NUMBER	DATE
PRIORITY INFORMATION:	US 1999-167035P	19991123 (60)
DOCUMENT TYPE:	Utility	
FILE SEGMENT:	GRANTED	
PRIMARY EXAMINER:	McGarry, Sean	
ASSISTANT EXAMINER:	Zara, Jane	
LEGAL REPRESENTATIVE:	Keown & Associates	
NUMBER OF CLAIMS:	34	
EXEMPLARY CLAIM:	1	
NUMBER OF DRAWINGS:	0 Drawing Figure(s); 0 Drawing Page(s)	
LINE COUNT:	3198	

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention relates to the inhibition of histone deacetylase. The invention provides compounds and methods for inhibiting histone deacetylase enzymatic activity. The invention also provides compositions and methods for treating cell proliferative diseases and conditions.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 342373-15-3P

(preparation of hydroxamic acids as inhibitors of histone deacetylase)

RN 342373-15-3 USPATFULL

CN 2-Furanpropanamide, tetrahydro-N-hydroxy-5-(2-oxo-2-phenylethyl)- (9CI)
(CA INDEX NAME)

